chromosome 8

Humans normally have 46 chromosomes in each cell, divided into 23 pairs. Two copies of chromosome 8, one copy inherited from each parent, form one of the pairs. Chromosome 8 spans more than 146 million DNA building blocks (base pairs) and represents between 4.5 and 5 percent of the total DNA in cells.

Identifying genes on each chromosome is an active area of genetic research. Because researchers use different approaches to predict the number of genes on each chromosome, the estimated number of genes varies. Chromosome 8 likely contains about 700 genes that provide instructions for making proteins. These proteins perform a variety of different roles in the body.

Health Conditions Related to Chromosomal Changes

The following chromosomal conditions are associated with changes in the structure or number of copies of chromosome 8.

8p11 myeloproliferative syndrome

Translocations of genetic material between chromosome 8 and other chromosomes can cause 8p11 myeloproliferative syndrome. This condition is characterized by an increased number of white blood cells (myeloproliferative disorder) and the development of lymphoma, a blood-related cancer that causes tumor formation in the lymph nodes. The myeloproliferative disorder usually develops into another form of blood cancer called acute myeloid leukemia. The most common translocation involved in this condition, written as t(8;13)(p11;q12), fuses part of the *FGFR1* gene on chromosome 8 with part of the *ZMYM2* gene on chromosome 13. The translocations are found only in cancer cells.

The protein produced from the normal *FGFR1* gene can turn on cellular signaling that helps the cell respond to its environment, for example by stimulating cell growth. The protein produced from the fused gene, regardless of the partner gene involved, leads to constant FGFR1 signaling. The uncontrolled signaling promotes continuous cell growth and division, leading to cancer.

core binding factor acute myeloid leukemia

A type of blood cancer known as core binding factor acute myeloid leukemia (CBF-AML) is associated with a rearrangement (translocation) of genetic material between chromosomes 8 and 21. This rearrangement is associated with approximately 7 percent of acute myeloid leukemia cases in adults. The translocation, written as t(8;21), fuses part of the *RUNX1T1* gene (also known as *ETO*) from chromosome 8

with part of the *RUNX1* gene from chromosome 21. This mutation is acquired during a person's lifetime and is present only in certain cells. This type of genetic change, called a somatic mutation, is not inherited.

The fusion protein produced from the t(8;21) translocation, called RUNX1-ETO, retains some function of the two individual proteins. The normal RUNX1 protein, produced from the *RUNX1* gene, is part of a protein complex called core binding factor (CBF) that attaches (binds) to DNA and turns on genes involved in blood cell development. The normal ETO protein, produced from the *RUNX1T1* gene, turns off gene activity. The fusion protein forms CBF and attaches to DNA, but instead of turning on genes that stimulate the development of blood cells, it turns those genes off. This change in gene activity blocks the maturation (differentiation) of blood cells and leads to the production of abnormal, immature white blood cells called myeloid blasts. While t(8;21) is important for leukemia development, one or more additional genetic changes are typically needed for the myeloid blasts to develop into cancerous leukemia cells.

Langer-Giedion syndrome

Langer-Giedion syndrome is caused by a deletion or mutation in several genes on the long (q) arm of chromosome 8 at a position described as 8q24.1. This condition causes bone abnormalities, including noncancerous bone tumors known as exostoses, and distinctive facial features. The signs and symptoms of this condition are related to the deletion or mutation in at least two genes from this part of the chromosome. Researchers have determined that the loss of a functional *EXT1* gene is responsible for the multiple noncancerous (benign) bone tumors called exostoses seen in people with Langer-Giedion syndrome. Loss of a functional *TRPS1* gene may cause the other bone and facial abnormalities. One copy of the *EXT1* gene and the *TRPS1* gene are always missing or mutated in affected individuals; however, neighboring genes may also be involved. The loss of additional genes from this region of chromosome 8 likely contributes to the varied features of Langer-Giedion syndrome.

recombinant 8 syndrome

A rearrangement of chromosome 8 causes recombinant 8 syndrome, a condition that involves heart and urinary tract abnormalities, moderate to severe intellectual disability, and a distinctive facial appearance. This rearrangement results in a deletion of a piece of the short (p) arm and a duplication of a piece of the long (q) arm. This chromosome abnormality is written rec(8)dup(8q)inv(8)(p23.1q22.1). The signs and symptoms of recombinant 8 syndrome are related to the loss of genetic material on the short arm of chromosome 8 and the presence of extra genetic material on the long arm of chromosome 8. Researchers are working to determine which genes are involved in the deletion and duplication on chromosome 8.

other cancers

Translocations between chromosome 8 and other chromosomes have been associated with other types of cancer. For example, Burkitt lymphoma (a cancer of white blood cells called B cells that occurs most often in children and young adults) can be caused by a translocation between chromosomes 8 and 14. This translocation, written t(8;14)(q24;q32), leads to continuous cell division without control or order, which likely contributes to the development of Burkitt lymphoma. Less frequently, Burkitt lymphoma can be caused by translocations between chromosomes 8 and 2 or chromosomes 8 and 22.

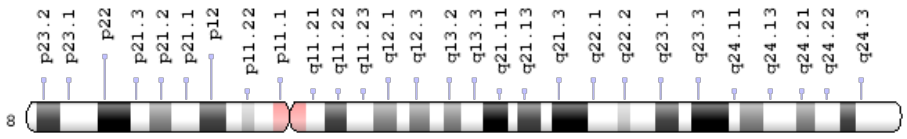
other chromosomal conditions

Trisomy 8 occurs when cells have three copies of chromosome 8 instead of the usual two copies. Full trisomy 8, which occurs when all of the body's cells contain an extra copy of chromosome 8, is not compatible with life. A similar but less severe condition called mosaic trisomy 8 occurs when only some of the body's cells have an extra copy of chromosome 8. The signs and symptoms of mosaic trisomy 8 vary widely and can include intellectual disability, absence of the tissue connecting the left and right halves of the brain (corpus callosum), skeletal defects, heart problems, kidney and liver malformations, and facial abnormalities. Trisomy 8 mosaicism is also associated with an increased risk of acute myeloid leukemia.

Another chromosomal condition called inversion duplication 8p is caused by a rearrangement of genetic material on the short (p) arm of chromosome 8. This rearrangement results in an abnormal duplication and an inversion of a segment of the chromosome. An inversion involves the breakage of a chromosome in two places; the resulting piece of DNA is reversed and reinserted into the chromosome. People with inversion duplication 8p typically have severe intellectual disability, a thin or absent corpus callosum, weak muscle tone (hypotonia), abnormal curvature of the spine (scoliosis), and minor facial abnormalities. Some individuals with this condition may also have heart defects, underdeveloped kidneys, or eye abnormalities. Older individuals usually develop abnormal muscle stiffness (spasticity). The signs and symptoms of inversion duplication 8p tend to depend on the size and location of the chromosome segment involved. For example, inclusion of chromosome region 8p21 is thought to be associated with more severe symptoms.

Chromosome Diagram

Geneticists use diagrams called idiograms as a standard representation for chromosomes. Idiograms show a chromosome's relative size and its banding pattern, which is the characteristic pattern of dark and light bands that appears when a chromosome is stained with a chemical solution and then viewed under a microscope. These bands are used to describe the location of genes on each chromosome.



Credit: Genome Decoration Page/NCBI

Additional Information & Resources

MedlinePlus

- Encyclopedia: Chromosome
  https://medlineplus.gov/ency/article/002327.htm

Additional NIH Resources

- National Human Genome Research Institute: Chromosome Abnormalities
  https://www.genome.gov/11508982/

Educational Resources

- Atlas of Genetics and Cytogenetics in Oncology and Haematology
  http://atlasgeneticsoncology.org/Anomalies/t0814ID1050.html

Scientific Articles on PubMed

- PubMed
  https://www.ncbi.nlm.nih.gov/pubmed?term=%28Chromosomes,+Human,+Pair+8%5BMAJR%5D%29+AND+%28Chromosome+8%5BTI%5D%29+AND+english%5Bla%5D+AND+human%5Bmh%5D+AND+%22last+1440+days%22%5Bdp%5D

OMIM

- BURKITT LYMPHOMA
  http://omim.org/entry/113970
- RECOMBINANT CHROMOSOME 8 SYNDROME
  http://omim.org/entry/179613
- TRICHORHINOPHALANGEAL SYNDROME, TYPE II
  http://omim.org/entry/150230

Sources for This Summary

- Ayakannu T, Wordsworth S, Parveen S, Shehadeh Z, Moselhi M. Rare presentation of Trisomy 8 syndrome. J Obstet Gynaecol. 2008 Oct;28(7):748-9. doi: 10.1080/01443610802462027.
  Citation on PubMed: https://www.ncbi.nlm.nih.gov/pubmed/19065376
- Ensembl Human Map View: Chromosome 8
  http://www.ensembl.org/Homo_sapiens/Location/Chromosome?chr=8;r=8:1-145138636
- Feenstra I, van Ravenswaaij CM, van der Knaap MS, Willemsen MA. Neuroimaging in nine patients with inversion duplication of the short arm of chromosome 8. Neuropediatrics. 2006 Apr;37(2):83-7.
  Citation on PubMed: https://www.ncbi.nlm.nih.gov/pubmed/16773506
- Gilbert F. Chromosome 8. Genet Test. 2001 Winter;5(4):345-54.
  Citation on PubMed: https://www.ncbi.nlm.nih.gov/pubmed/11960583
- Golzio C, Guirchoun J, Ozilou C, Thomas S, Goudefroye G, Morichon-Delvallez N, Vekemans M, Attié-Bitach T, Etchevers HC. Cytogenetic and histological features of a human embryo with homogeneous chromosome 8 trisomy. Prenat Diagn. 2006 Dec;26(13):1201-5.
  Citation on PubMed: https://www.ncbi.nlm.nih.gov/pubmed/17075794
- Graw SL, Sample T, Bleskan J, Sujansky E, Patterson D. Cloning, sequencing, and analysis of inv8 chromosome breakpoints associated with recombinant 8 syndrome. Am J Hum Genet. 2000 Mar; 66(3):1138-44.
  Citation on PubMed: https://www.ncbi.nlm.nih.gov/pubmed/10712224
  Free article on PubMed Central: https://www.ncbi.nlm.nih.gov/pmc/articles/PMC1288148/
- Hulley BJ, Hummel M, Cook LL, Boyd BK, Wenger SL. Trisomy 8 mosaicism: selective growth advantage of normal cells vs. growth disadvantage of trisomy 8 cells. Am J Med Genet A. 2003 Jan 15;116A(2):144-6.
  Citation on PubMed: https://www.ncbi.nlm.nih.gov/pubmed/12494432
- Jackson CC, Medeiros LJ, Miranda RN. 8p11 myeloproliferative syndrome: a review. Hum Pathol. 2010 Apr;41(4):461-76. doi: 10.1016/j.humpath.2009.11.003. Review.
  Citation on PubMed: https://www.ncbi.nlm.nih.gov/pubmed/20226962
- Lam K, Zhang DE. RUNX1 and RUNX1-ETO: roles in hematopoiesis and leukemogenesis. Front Biosci (Landmark Ed). 2012 Jan 1;17:1120-39. Review.
  Citation on PubMed: https://www.ncbi.nlm.nih.gov/pubmed/22201794
  Free article on PubMed Central: https://www.ncbi.nlm.nih.gov/pmc/articles/PMC3433167/
- Map Viewer: Genes on Sequence
  https://www.ncbi.nlm.nih.gov/mapview/maps.cgi?ORG=human&MAPS=ideogr,ugHs,genes&CHR=8

- Merchant S, Schlette E, Sanger W, Lai R, Medeiros LJ. Mature B-cell leukemias with more than 55% prolymphocytes: report of 2 cases with Burkitt lymphoma-type chromosomal translocations involving c-myc. Arch Pathol Lab Med. 2003 Mar;127(3):305-9. Review.
  Citation on PubMed: https://www.ncbi.nlm.nih.gov/pubmed/12653573
- Nusbaum C, Mikkelsen TS, Zody MC, Asakawa S, Taudien S, Garber M, Kodira CD, Schueler MG, Shimizu A, Whittaker CA, Chang JL, Cuomo CA, Dewar K, FitzGerald MG, Yang X, Allen NR, Anderson S, Asakawa T, Blechschmidt K, Bloom T, Borowsky ML, Butler J, Cook A, Corum B, DeArellano K, DeCaprio D, Dooley KT, Dorris L 3rd, Engels R, Glöckner G, Hafez N, Hagopian DS, Hall JL, Ishikawa SK, Jaffe DB, Kamat A, Kudoh J, Lehmann R, Lokitsang T, Macdonald P, Major JE, Matthews CD, Mauceli E, Menzel U, Mihalev AH, Minoshima S, Murayama Y, Naylor JW, Nicol R, Nguyen C, O'Leary SB, O'Neill K, Parker SC, Polley A, Raymond CK, Reichwald K, Rodriguez J, Sasaki T, Schilhabel M, Siddiqui R, Smith CL, Sneddon TP, Talamas JA, Tenzin P, Topham K, Venkataraman V, Wen G, Yamazaki S, Young SK, Zeng Q, Zimmer AR, Rosenthal A, Birren BW, Platzer M, Shimizu N, Lander ES. DNA sequence and analysis of human chromosome 8. Nature. 2006 Jan 19;439(7074):331-5.
  Citation on PubMed: https://www.ncbi.nlm.nih.gov/pubmed/16421571
- Pienkowska-Grela B, Witkowska A, Grygalewicz B, Rymkiewicz G, Rygier J, Woroniecka R, Walewski J. Frequent aberrations of chromosome 8 in aggressive B-cell non-Hodgkin lymphoma. Cancer Genet Cytogenet. 2005 Jan 15;156(2):114-21.
  Citation on PubMed: https://www.ncbi.nlm.nih.gov/pubmed/15642390
- Smith AC, Spuhler K, Williams TM, McConnell T, Sujansky E, Robinson A. Genetic risk for recombinant 8 syndrome and the transmission rate of balanced inversion 8 in the Hispanic population of the southwestern United States. Am J Hum Genet. 1987 Dec;41(6):1083-103.
  Citation on PubMed: https://www.ncbi.nlm.nih.gov/pubmed/3687942
  Free article on PubMed Central: https://www.ncbi.nlm.nih.gov/pmc/articles/PMC1684361/
- UCSC Genome Browser: Statistics
  http://genome.cse.ucsc.edu/goldenPath/stats.html
- Voigt R, Gburek-Augustat J, Seidel A, Gillessen-Kaesbach G. Hemihyperplasia and discordant bone age in a patient with trisomy 8 mosaicism. Am J Med Genet A. 2008 Jan 1;146A(1):132-5.
  Citation on PubMed: https://www.ncbi.nlm.nih.gov/pubmed/18074372
- Xiao S, Nalabolu SR, Aster JC, Ma J, Abruzzo L, Jaffe ES, Stone R, Weissman SM, Hudson TJ, Fletcher JA. FGFR1 is fused with a novel zinc-finger gene, ZNF198, in the t(8;13) leukaemia/ lymphoma syndrome. Nat Genet. 1998 Jan;18(1):84-7.
  Citation on PubMed: https://www.ncbi.nlm.nih.gov/pubmed/9425908
- de Die-Smulders CE, Engelen JJ, Schrander-Stumpel CT, Govaerts LC, de Vries B, Vles JS, Wagemans A, Schijns-Fleuren S, Gillessen-Kaesbach G, Fryns JP. Inversion duplication of the short arm of chromosome 8: clinical data on seven patients and review of the literature. Am J Med Genet. 1995 Nov 20;59(3):369-74. Review.
  Citation on PubMed: https://www.ncbi.nlm.nih.gov/pubmed/8599364

Reprinted from Genetics Home Reference:
https://ghr.nlm.nih.gov/chromosome/8.pdf

Reviewed: November 2013
Published: March 21, 2017

Lister Hill National Center for Biomedical Communications
U.S. National Library of Medicine
National Institutes of Health
Department of Health & Human Services